IAP20 Rec'd PCT/PTO 23 JUN 2006

SEQUENCE LISTING

| <110> KIRIN BEER KABUSIKI .KAISHA |
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| <120> A mutant of anti CD40 antibody |
| <130> PH-2356-PCT |
| <140> <141> |
| <150> JP 2003-431408 <151> 2003-12-25 |
| <160> 142 |
| <170> PatentIn Ver. 2.1 |
| <210> 1 |
| <211> 175 |
| <212> PRT |
| <213> Homo sapiens |
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| 1 5 10 15 |
| Cys Cys Ser Leu Cys Gln Pro Gly Gln Lys Leu Val Ser Asp Cys Thr |

25

30

Glu Phe Thr Glu Thr Glu Cys Leu Pro Cys Gly Glu Ser Glu Phe Leu
35 40 45

Asp Thr Trp Asn Arg Glu Thr His Cys His Gln His Lys Tyr Cys Asp
50 55 60

Pro Asn Leu Gly Leu Arg Val Gln Gln Lys Gly Thr Ser Glu Thr Asp
65 70 75 80

Thr Ile Cys Thr Cys Glu Glu Gly Trp His Cys Thr Ser Glu Ala Cys
85 90 95

Glu Ser Cys Val Leu His Arg Ser Cys Ser Pro Gly Phe Gly Val Lys
100 105 110

Gln Ile Ala Thr Gly Val Ser Asp Thr Ile Cys Glu Pro Cys Pro Val 115 120 125

Gly Phe Phe Ser Asn Val Ser Ser Ala Phe Glu Lys Cys His Pro Trp
130 135 140

Thr Ser Cys Glu Thr Lys Asp Leu Val Val Gln Gln Ala Gly Thr Asn 145 150 155 160

Lys Thr Asp Val Val Cys Gly Pro Gln Asp Arg Leu Arg Ala Leu
165 170 175

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| ⟨220⟩ |
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| ⟨400⟩ 2 |
| atatgctagc accaagggcc catcggtctt cccctggc 39 |
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| ⟨210⟩ 3 |
| ⟨211⟩ 38 |
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40

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<213> Artificial Sequence

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26

<210> 6

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<213> Artificial Sequence

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<223> Description of Artificial Sequence: Synthetic DNA

⟨400⟩ 6

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38

<210> 7

| <212> DNA | |
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| ⟨210⟩ 8 | |
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aggggtccgg gagatcatga gagtgtcctt

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<210> 10

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<211> 23

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<400> 11

tgatcatacg tagatatcac ggc

23

<210> 12

| <212> DNA | |
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| ⟨220⟩ | |
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| ⟨400⟩ 12 | |
| tgatcatacg tagatatcac ggc | 23 |
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| <400> 13 | |
| gggtacgtcc tcacattcag tgatcag | 27 |
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| <211> 39 | |
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| <220> | |
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| <210> 15 |
| <211> 23 |
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<210> 17<211> 27

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| <213> Artificial Sequence | |
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| <400> 17 | |
| gggtacgtcc tcacattcag tgatcag | 27 |
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| <210> 18 | |
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| <212> DNA | |
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| <400> 18 | |
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| <210> 19 | |
| <211> 31 | |
| <212> DNA | |
| <213> Artificial Sequence | |
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| <400> | 19 |
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| <210> | 20 |
| <211> | 23 |
| <212> | DNA |
| <213> | Artificial Sequence |
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| <220> | |
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| <400> | 20 |
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| <210> | 21 |
| <211> | 27 |
| <212> | DNA |
| <213> | Artificial Sequence |
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| <220> | |
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27

<210> 22 <211> 31

<400> 21

gggtacgtcc tcacattcag tgatcag

| <212> DNA | |
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| <213> Artificial Sequence | |
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| ⟨220⟩ | |
| <223> Description of Artificial Sequence: Synthetic DNA | |
| | |
| <400> 22 | |
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| <210> 23 | |
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| <212> DNA | |
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| | |
| <220> | |
| <223> Description of Artificial Sequence:Synthetic DNA | |
| • | |
| ⟨400⟩ 23 | |
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| | |
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| <212> DNA | |
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| <220> | |
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| ⟨220⟩ | |
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| <210> 26 | |
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| <400> 26 | |
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<400> 24

⟨210⟩ 27

⟨211⟩ 32

| <212> DNA | ٠ |
|---|----|
| <213> Artificial Sequence | |
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| <223> Description of Artificial Sequence:Synthetic DNA | |
| | |
| <400> 27 | |
| ggtggacaag agagttgagt ccaaatgttg tg | 32 |
| | |
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| <210> 28 | |
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| <220> | |
| <pre><223> Description of Artificial Sequence:Synthetic DNA</pre> | |
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| tgatcatacg tagatatcac ggc | 23 |
| | |
| Z010\ 00 | |
| <210> 29 | |
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| <212> DNA | |
| <213> Artificial Sequence | |
| /990 \ | |
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| <pre><223> Description of Artificial Sequence:Synthetic DNA</pre> | |

| 110 | ^ | 29 |
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| <40 | U2 | 7.9 |

gggtacgtcc tcacattcag tgatcag

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<211> 34

<212> DNA

<213> Artificial Sequence

<220>

<223> Description of Artificial Sequence: Synthetic DNA

<400> 30

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<210> 31

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<213> Artificial Sequence

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<400> 31

gagcgcaaat atggtccccc atgcccaccg tgcc

34

<210> 32

| <212> DNA | |
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| ⟨220⟩ | |
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| ⟨400⟩ 32 | |
| tgatcatacg tagatatcac ggc | 23 |
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| | |
| ⟨210⟩ 33 | |
| <211> 27 | |
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| <220> | |
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| , | • |
| <400> 33 | |
| gggtacgtcc tcacattcag tgatcag | 27 |
| | |
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| ⟨210⟩ 34 | |
| ⟨211⟩ 39 | |
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| <220> | |
| <223> Description of Artificial Sequence: Synthet | cic DNA |

| <400> | 34 | | | |
|--|--|--|--|--|
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| | | | | |
| <210> | 35 | | | |
| <211> | 39 | | | |
| <212> | DNA | | | |
| <213> | Artificial Sequence | | | |
| | • | | | |
| <220> | | | | |
| <223> | Description of Artificial Sequence:Synthetic DNA | | | |
| | | | | |
| <400> | 35 | | | |
| tgcccagcac cagagttcct ggggggaccg tcagtcttc | | | | |
| | • • | | | |
| | | | | |
| <210> | 36 | | | |
| <211> | 23 | | | |
| <212> | DNA | | | |
| <213> | Artificial Sequence | | | |
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39

<400> 36
tgatcatacg tagatatcac ggc 23

 $\ensuremath{\texttt{<}223\texttt{>}}$ Description of Artificial Sequence:Synthetic DNA

<210> 37 <211> 1480

<212> DNA

<213> Homo sapiens

<400> 37

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| Tyr Pro Tyr 130 | Tyr Ty | | Met Asp 135 | Val T | rp Gly | Gln Gly 140 | Thr T | hr Val |
|--------------------|----------------|---------|----------------|--------------|----------------|----------------|---------|---------------|
| Thr Val Ser | Ser Ala | 150 Ser | Γhr Lys | Gly P | Pro Ser 155 | Val Phe | e Pro L | eu Ala 160 |
| Pro Cys Ser | Arg Se | | Ser Glu | | Chr Ala | Ala Leu | | ys Leu 75 |
| Val Lys Asp | Tyr Pho | e Pro (| Glu Pro | Val T | ìhr Val | Ser Trp | Asn S | er Gly |
| Ala Leu Thr 195 | Ser Gl | v Val H | His Thr 200 | Phe P | Pro Ala | Val Leu 209 | | er Ser |
| Gly Leu Tyr 210 | Ser Le | | Ser Val 215 | Val T | Chr Val | Pro Sei | Ser A | sn Phe |
| Gly Thr Gln 225 | Thr Ty | Thr (| Cys Asn | Val A | Asp His 235 | Lys Pro | Ser A | sn Thr 240 |
| Lys Val Asp | Lys Th | | Glu Arg | | Cys Cys 250 | Val Glu | | ro Pro 55 |
| Cys Pro Ala | Pro Pro 260 | o Val A | Ala Gly | Pro S 265 | Ser Val | Phe Let | Phe P | ro Pro |
| Lys Pro Lys | Asp Th | Leu M | Met Ile | Ser A | arg Thr | Pro Glu | ı Val T | hr Cys |

Val Val Val Asp Val Ser His Glu Asp Pro Glu Val Gln Phe Asn Trp
290
295
300

Tyr Val Asp Gly Val Glu Val His Asn Ala Lys Thr Lys Pro Arg Glu 305 310 315 320

Glu Gln Phe Asn Ser Thr Phe Arg Val Val Ser Val Leu Thr Val Val
325
330
335

His Gln Asp Trp Leu Asn Gly Lys Glu Tyr Lys Cys Lys Val Ser Asn 340 345 350

Lys Gly Leu Pro Ala Pro Ile Glu Lys Thr Ile Ser Lys Thr Lys Gly
355 360 365

Gln Pro Arg Glu Pro Gln Val Tyr Thr Leu Pro Pro Ser Arg Glu Glu 370 375 380

Met Thr Lys Asn Gln Val Ser Leu Thr Cys Leu Val Lys Gly Phe Tyr 385 390 395 400

Pro Ser Asp Ile Ala Val Glu Trp Glu Ser Asn Gly Gln Pro Glu Asn
405
410
415

Asn Tyr Lys Thr Thr Pro Pro Met Leu Asp Ser Asp Gly Ser Phe Phe
420 425 430

Leu Tyr Ser Lys Leu Thr Val Asp Lys Ser Arg Trp Gln Gln Gly Asn
435
440
445

Val Phe Ser Cys Ser Val Met His Glu Ala Leu His Asn His Tyr Thr 450 455 460

Gln Lys Ser Leu Ser Leu Ser Pro Gly Lys
465 470

<210> 39

<211> 406

<212> DNA

<213> Homo sapiens

<400> 39

actgctcagt taggacccag agggaaccat ggaagcccca gctcagcttc tettectect 60 gctactctgg ctcccagata ccaccggaga aattgtgttg acacagtetc cagccaccet 120 gtctttgtet ccaggggaaa gagccaccet etcetgcagg gccagtcaga gtgttagcag 180 ctacttagce tggtaccaac agaaacctgg ccaggetece aggetectca tetatgatge 240 atccaacagg gccactggca tcccagccag gttcagtggc agtgggtetg ggacagactt 300 cactetcace atcagcagce tagagcetga agattttgca gtttattact gtcagcagcg 360 tagcaacact ttcggccetg ggaccaaagt ggatatcaaa cgtacg 406

<210> 40

<211> 126

<212> PRT

<213> Homo sapiens

<400> 40

Met Glu Ala Pro Ala Gln Leu Leu Phe Leu Leu Leu Leu Trp Leu Pro

10

5

1

15

Asp Thr Thr Gly Glu Ile Val Leu Thr Gln Ser Pro Ala Thr Leu Ser
20 25 30

Leu Ser Pro Gly Glu Arg Ala Thr Leu Ser Cys Arg Ala Ser Gln Ser

35 40 45

Val Ser Ser Tyr Leu Ala Trp Tyr Gln Gln Lys Pro Gly Gln Ala Pro
50 55 60

Arg Leu Leu Ile Tyr Asp Ala Ser Asn Arg Ala Thr Gly Ile Pro Ala 65 70 75 80

Arg Phe Ser Gly Ser Gly Ser Gly Thr Asp Phe Thr Leu Thr Ile Ser 85 90 95

Ser Leu Glu Pro Glu Asp Phe Ala Val Tyr Tyr Cys Gln Gln Arg Ser 100 105 110

Asn Thr Phe Gly Pro Gly Thr Lys Val Asp Ile Lys Arg Thr
115 120 125

<210> 41

<211> 508

<212> DNA

<213> Homo sapiens

<400> 41

ctgaacacag acccgtcgac tcccaggtgt ttccattcag tgatcagcac tgaacacaga 60 ggactcacca tggagttggg actgagctgg attttccttt tggctatttt aaaaggtgtc 120 cagtgtgaag tgcagctggt ggagtctggg ggaggcttgg tacagcctgg caggtccctg 180 agactctcct gtgcagcctc tggattcacc tttgatgatt atgccatgca ctgggtccgg 240 caagctccag ggaagggcct ggagtggtc tcaggtatta gttggaatag tggtagcttg 300 gtgcatgcgg actctgtgaa gggccgattc accatctcca gagacaacgc caagaactcc 360 ctgtatctgc aaatgaacag tctgagagct gaggacacgg ccttgtatta ctgtgcaaga 420 gataggctat ttcgggagt taggtactac ggtatggacg tctggggcca agggaccacg 480 gtcaccgtct cctcagctag caccaagg

<210> 42

<211> 146

<212> PRT

<213> Homo sapiens

<400> 42

Met Glu Leu Gly Leu Ser Trp Ile Phe Leu Leu Ala Ile Leu Lys Gly

1 5 10 15

Val Gln Cys Glu Val Gln Leu Val Glu Ser Gly Gly Gly Leu Val Gln
20 25 30

Pro Gly Arg Ser Leu Arg Leu Ser Cys Ala Ala Ser Gly Phe Thr Phe
35 40 45

Asp Asp Tyr Ala Met His Trp Val Arg Gln Ala Pro Gly Lys Gly Leu 50 55 60

Glu Trp Val Ser Gly Ile Ser Trp Asn Ser Gly Ser Leu Val His Ala

65 70 75

80

Asp Ser Val Lys Gly Arg Phe Thr Ile Ser Arg Asp Asn Ala Lys Asn 85 90 95

Ser Leu Tyr Leu Gln Met Asn Ser Leu Arg Ala Glu Asp Thr Ala Leu 100 105 110

Tyr Tyr Cys Ala Arg Asp Arg Leu Phe Arg Gly Val Arg Tyr Tyr Gly
115 120 125

Met Asp Val Trp Gly Gln Gly Thr Thr Val Thr Val Ser Ser Ala Ser
130 135 140

Thr Lys

145

<210> 43

<211> 414

<212> DNA

<213> Homo sapiens

<400> 43

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<210> 44

<211> 129

<212> PRT

<213> Homo sapiens

<400> 44

Met Glu Ala Pro Ala Gln Leu Leu Phe Leu Leu Leu Leu Trp Leu Pro

1 5 10 15

Asp Thr Thr Gly Glu Ile Val Leu Thr Gln Ser Pro Ala Thr Leu Ser
20 25 30

Leu Ser Pro Gly Glu Arg Ala Thr Leu Ser Cys Arg Ala Ser Gln Ser
35 40 45

Val Ser Ser Tyr Leu Ala Trp Tyr Gln Gln Lys Pro Gly Gln Ala Pro
50 55 60

Arg Leu Leu Ile Tyr Asp Ala Ser Asn Arg Ala Thr Gly Ile Pro Ala 65 70 75 80

Arg Phe Ser Gly Ser Gly Ser Gly Thr Asp Phe Thr Leu Thr Ile Ser 85 90 95

Ser Leu Glu Pro Glu Asp Phe Ala Val Tyr Tyr Cys Gln Gln Arg Ser 100 105 110

His Trp Leu Thr Phe Gly Gly Gly Thr Lys Val Glu Ile Lys Arg Thr 115 120 125

Val

<210> 45

<211> 462

<212> DNA

<213> Homo sapiens

<400> 45

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<210> 46

<211> 149

<212> PRT

<213> Homo sapiens

<400> 46

 $\hbox{Met Asp Leu Met Cys Lys Lys Met Lys His Leu Trp\ Phe\ Phe\ Leu\ Leu}$

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Leu Val Ala Ala Pro Arg Trp Val Leu Ser Gln Leu Gln Leu Gln Glu
20 25 30

Ser Gly Pro Gly Leu Leu Lys Pro Ser Glu Thr Leu Ser Leu Thr Cys

35 40 45

Thr Val Ser Gly Gly Ser Ile Ser Ser Pro Gly Tyr Tyr Gly Gly Trp
50 55 60

Ile Arg Gln Pro Pro Gly Lys Gly Leu Glu Trp Ile Gly Ser Ile Tyr
65 70 75 80

Lys Ser Gly Ser Thr Tyr His Asn Pro Ser Leu Lys Ser Arg Val Thr
85 90 95

Ile Ser Val Asp Thr Ser Lys Asn Gln Phe Ser Leu Lys Leu Ser Ser

100 105 110

Val Thr Ala Ala Asp Thr Ala Val Tyr Tyr Cys Thr Arg Pro Val Val
115 120 125

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| Tyr Pro Tyr 130 | Tyr Tyr | Ser Met | Asp Val | Trp Gly | Gln Gly 140 | Thr Thr Val |
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| Pro Cys Ser | Arg Ser 165 | Thr Ser | Glu Ser | Thr Ala | Ala Leu | Gly Cys Leu 175 |
| Val Lys Asp | Tyr Phe | Pro Glu | Pro Val | | Ser Trp | Asn Ser Gly |
| Ala Leu Thr 195 | Ser Gly | Val His | Thr Phe | Pro Ala | Val Leu 205 | Gln Ser Ser |
| Gly Leu Tyr 210 | Ser Leu | Ser Ser 215 | Val Val | Thr Val | Pro Ser 220 | Ser Asn Phe |
| Gly Thr Gln 225 | Thr Tyr | Thr Cys | Asn Val | Asp His | Lys Pro | Ser Asn Thr 240 |
| Lys Val Asp | Lys Thr | Val Glu | Arg Lys | Cys Cys 250 | Val Glu | Cys Pro Pro 255 |
| Cys Pro Ala | Pro Pro 260 | Val Ala | Gly Pro | | Phe Leu | Phe Pro Pro 270 |
| Lys Pro Lys 275 | Asp Thr | Leu Met | Ile Ser 280 | Arg Thr | Pro Glu 285 | Val Thr Cys |

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<213> Homo sapiens

<400> 133

atggaagccc cagctcagct tetetteete etgetaetet ggeteeraga taccacegga 60 gaaattgtgt tgacacagte tecagecaec etgetettgt etecagegga aagagecaec 120 eteteetgea gggecagtea gagtgtage agetaettag eetggaaca acagaaacet 180 ggecaggete eateetatgat geateeraaca gggecaetgg eateecagee 240 aggtteagtg geagtggte tgggacagae tteaetetea eeateageag eetaggaeet 300 gaagattttg eagttatta etgeteageag egtageaaca ettteegeee tgggaceaaa 360 gtggatatea aacgtaeggt ggetgeaeca tetgtettea tetteegeee atetgatgag 420 eagttgaaat etggaactge etetgttgt tgeetgetga ataaetteta teecagagag 480 gecaaagtae agtggaaggt ggataacgee eteeaategg gtaaeteeea ggagagtgte 540 acagageagg acagcaagaa eageaectae ageeteagaa geaecetgae getgageaaa 600 geagaetaec agaacaaa agtetaegee tgegaagtea eecaateagg eetgageaa 660 eeegteaaaa agagetteaa eaggggagag tgttga

<210> 134

<211> 231

<212> PRT

<213> Homo sapiens

<400> 134

Met Glu Ala Pro Ala Gln Leu Leu Phe Leu Leu Leu Leu Trp Leu Pro

1 5 10 15

Asp Thr Thr Gly Glu Ile Val Leu Thr Gln Ser Pro Ala Thr Leu Ser
20 25 30

Leu Ser Pro Gly Glu Arg Ala Thr Leu Ser Cys Arg Ala Ser Gln Ser 35 40 45

Val Ser Ser Tyr Leu Ala Trp Tyr Gln Gln Lys Pro Gly Gln Ala Pro
50 55 60

Arg Leu Leu Ile Tyr Asp Ala Ser Asn Arg Ala Thr Gly Ile Pro Ala 65 70 75 80

Arg Phe Ser Gly Ser Gly Thr Asp Phe Thr Leu Thr Ile Ser

85 90 95

Ser Leu Glu Pro Glu Asp Phe Ala Val Tyr Tyr Cys Gln Gln Arg Ser

100 105 110

Asn Thr Phe Gly Pro Gly Thr Lys Val Asp Ile Lys Arg Thr Val Ala 115 120 125

Ala Pro Ser Val Phe Ile Phe Pro Pro Ser Asp Glu Gln Leu Lys Ser 130 135 140 Gly Thr Ala Ser Val Val Cys Leu Leu Asn Asn Phe Tyr Pro Arg Glu

145 150 155 160

Ala Lys Val Gln Trp Lys Val Asp Asn Ala Leu Gln Ser Gly Asn Ser

165 170 175

Gln Glu Ser Val Thr Glu Gln Asp Ser Lys Asp Ser Thr Tyr Ser Leu 180 185 190

Ser Ser Thr Leu Thr Leu Ser Lys Ala Asp Tyr Glu Lys His Lys Val
195 200 205

Tyr Ala Cys Glu Val Thr His Gln Gly Leu Ser Ser Pro Val Thr Lys 210 215 220

Ser Phe Asn Arg Gly Glu Cys

225 230

<210> 135

<211> 1407

<212> DNA

<213> Homo sapiens

<400> 135

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gactetgtga agggeegatt caccatetee agagacaaeg ceaagaaete eetgtatetg 300 caaatgaaca gtctgagagc tgaggacacg gccttgtatt actgtgcaag agataggcta 360 tttcggggag ttaggtacta cggtatggac gtctggggcc aagggaccac ggtcaccgtc 420 tecteageta geaceaaggg eccateggte tteceetgg egeeetgete eaggageace 480 tecgagagea cageggeeet gggetgeetg gteaaggaet actteeeega aceggtgaeg 540 gtgtcgtgga actcaggcgc tctgaccagc ggcgtgcaca ccttcccagc tgtcctacag 600 tectcaggae tetacteect cagcagegtg gtgacegtge cetecageaa etteggeace 660 cagacetaca cetgeaacgt agateacaag eccageaaca ceaaggtgga caagacagtt 720 gagcgcaaat gttgtgtcga gtgcccaccg tgcccagcac cacctgtggc aggaccgtca 780 gtetteetet teeceecaaa acceaaggae acceteatga teteeeggae eeetgaggte 840 acgtgcgtgg tggtggacgt gagccacgaa gaccccgagg tccagttcaa ctggtacgtg 900 gacggcgtgg aggtgcataa tgccaagaca aagccacggg aggagcagtt caacagcacg 960 ttccgtgtgg tcagcgtcct caccgttgtg caccaggact ggctgaacgg caaggagtac 1020 aagtgcaagg tetecaacaa aggeeteeca geeteeateg agaaaaceat etecaaaace 1080 aaagggcagc cccgagaacc acaggtgtac accctgcccc catcccggga ggagatgacc 1140 aagaaccagg tcagcctgac ctgcctggtc aaaggcttct accccagcga catcgccgtg 1200 gagtgggaga gcaatgggca gccggagaac aactacaaga ccacacctcc catgctggac 1260 tecgaegget cettetteet etacageaag eteacegtgg acaagageag gtggeageag 1320 gggaacgtet teteatgete egtgatgeat gaggetetge acaaccacta caegeagaag 1380 agcctctccc tgtctccggg taaatga. 1407

<210> 136

<211> 468

<212> PRT

<213> Homo sapiens

<400> 136

Met Glu Leu Gly Leu Ser Trp Ile Phe Leu Leu Ala Ile Leu Lys Gly

1

5

10

15

Val Gln Cys Glu Val Gln Leu Val Glu Ser Gly Gly Leu Val Gln Pro Gly Arg Ser Leu Arg Leu Ser Cys Ala Ala Ser Gly Phe Thr Phe Asp Asp Tyr Ala Met His Trp Val Arg Gln Ala Pro Gly Lys Gly Leu Glu Trp Val Ser Gly Ile Ser Trp Asn Ser Gly Ser Leu Val His Ala Asp Ser Val Lys Gly Arg Phe Thr Ile Ser Arg Asp Asn Ala Lys Asn Ser Leu Tyr Leu Gln Met Asn Ser Leu Arg Ala Glu Asp Thr Ala Leu Tyr Tyr Cys Ala Arg Asp Arg Leu Phe Arg Gly Val Arg Tyr Tyr Gly Met Asp Val Trp Gly Gln Gly Thr Thr Val Thr Val Ser Ser Ala Ser Thr Lys Gly Pro Ser Val Phe Pro Leu Ala Pro Cys Ser Arg Ser Thr Ser Glu Ser Thr Ala Ala Leu Gly Cys Leu Val Lys Asp Tyr Phe Pro

Glu Pro Val Thr Val Ser Trp Asn Ser Gly Ala Leu Thr Ser Gly Val

180 185 190

His Thr Phe Pro Ala Val Leu Gln Ser Ser Gly Leu Tyr Ser Leu Ser 195 200 205

Ser Val Val Thr Val Pro Ser Ser Asn Phe Gly Thr Gln Thr Tyr Thr 210 215 220

Cys Asn Val Asp His Lys Pro Ser Asn Thr Lys Val Asp Lys Thr Val 225 230 235 240

Glu Arg Lys Cys Cys Val Glu Cys Pro Pro Cys Pro Ala Pro Pro Val
245 250 255

Ala Gly Pro Ser Val Phe Leu Phe Pro Pro Lys Pro Lys Asp Thr Leu 260 265 270

Met Ile Ser Arg Thr Pro Glu Val Thr Cys Val Val Val Asp Val Ser 275 280 285

His Glu Asp Pro Glu Val Gln Phe Asn Trp Tyr Val Asp Gly Val Glu 290 295 300

Val His Asn Ala Lys Thr Lys Pro Arg Glu Glu Gln Phe Asn Ser Thr 305 310 315 320

Phe Arg Val Val Ser Val Leu Thr Val Val His Gln Asp Trp Leu Asn 77/88

325 330 335

Gly Lys Glu Tyr Lys Cys Lys Val Ser Asn Lys Gly Leu Pro Ala Ser 340 345 350

Ile Glu Lys Thr Ile Ser Lys Thr Lys Gly Gln Pro Arg Glu Pro Gln
355 360 365

Val Tyr Thr Leu Pro Pro Ser Arg Glu Glu Met Thr Lys Asn Gln Val 370 375 380

Ser Leu Thr Cys Leu Val Lys Gly Phe Tyr Pro Ser Asp Ile Ala Val 385 390 395 400

Glu Trp Glu Ser Asn Gly Gln Pro Glu Asn Asn Tyr Lys Thr Thr Pro
405 410 415

Pro Met Leu Asp Ser Asp Gly Ser Phe Phe Leu Tyr Ser Lys Leu Thr
420 425 430

Val Asp Lys Ser Arg Trp Gln Gln Gly Asn Val Phe Ser Cys Ser Val
435
440
445

Met His Glu Ala Leu His Asn His Tyr Thr Gln Lys Ser Leu Ser Leu 450 455 460

Ser Pro Gly Lys

465

⟨210⟩ 137

<211> 702

<212> DNA

<213> Homo sapiens

<400> 137

atggaagece cageteaget tetetteete etgetaetet ggeteceaga taceacegga 60 gaaattgtgt tgacacagte tecagecace etgetettgt etceaggga aagagecace 120 eteteetgea gggecagtea gagtgttage agetaettag eetggtaeca acagaaacet 180 ggecaggete ecaggeteet catetatgat geatecaaca gggecaetgg cateceagee 240 aggtteagtg geagtgggte tgggacagae tteaetetea ecateageag eetaggget 300 gaagattttg eagtttatta etgteageag egtagecaet ggeteaettt eggegggggg 360 aceaaggtgg agateaaacg tacggtgget geaceatetg tetteatett eeggeggggg 360 aceaaggtgg tgaaatetgg aactgeetet gttgtgtgee tgetgaataa ettetatee 420 gatgageagt tgaaatetgg aactgeetet gttgtgtgee tgetgaataa ettetateee 480 agagagggea aagtacagg gaaggtggat aaegeeetee aategggtaa eteecaggag 540 agtgteacag ageaggaag caaggacage acetaeagee teageageae eetgaggetg 600 ageaaageag actaegagaa acacaaagte tacgeetgeg aagteacea teagggeetg 660 agetegeeeg teacaaagag etteaacagg ggagagtgtt ga 702

<210> 138

<211> 233

<212> PRT

<213> Homo sapiens

<400> 138

Met Glu Ala Pro Ala Gln Leu Leu Phe Leu Leu Leu Leu Trp Leu Pro

1 5 10 15

Asp Thr Thr Gly Glu Ile Val Leu Thr Gln Ser Pro Ala Thr Leu Ser 79/88

| 20 | 25 | 30 |
|----|----|----|
| | | |

| Leu | Ser | Pro | Gly | Glu | Arg | Ala | Thr | Leu | Ser | Cys | Arg | Ala | Ser | Gln | Ser |
|-----|-----|-----|-----|-----|-----|-----|-----|-----|-----|-----|-----|-----|-----|-----|-----|
| | | 35 | | | | | 40 | | | | | 45 | | | |

Val Ser Ser Tyr Leu Ala Trp Tyr Gln Gln Lys Pro Gly Gln Ala Pro
50 55 60

Arg Leu Leu Ile Tyr Asp Ala Ser Asn Arg Ala Thr Gly Ile Pro Ala 65 70 75 80

Arg Phe Ser Gly Ser Gly Thr Asp Phe Thr Leu Thr Ile Ser 85 90 95

Ser Leu Glu Pro Glu Asp Phe Ala Val Tyr Tyr Cys Gln Gln Arg Ser 100 105 110

His Trp Leu Thr Phe Gly Gly Gly Thr Lys Val Glu Ile Lys Arg Thr
115 120 125

Val Ala Ala Pro Ser Val Phe Ile Phe Pro Pro Ser Asp Glu Gln Leu
130 135 140

Lys Ser Gly Thr Ala Ser Val Val Cys Leu Leu Asn Asn Phe Tyr Pro 145 150 155 160

Arg Glu Ala Lys Val Gln Trp Lys Val Asp Asn Ala Leu Gln Ser Gly

165 170 175

Asn Ser Glìn Glu Ser Val Thr Glu Gln Asp Ser Lys Asp Ser Thr Tyr 80/88 180 185 190

Ser Leu Ser Ser Thr Leu Thr Leu Ser Lys Ala Asp Tyr Glu Lys His
195 200 205

Lys Val Tyr Ala Cys Glu Val Thr His Gln Gly Leu Ser Ser Pro Val 210 215 220

Thr Lys Ser Phe Asn Arg Gly Glu Cys 225 230

⟨210⟩ 139

<211> 1425

<212> DNA

<213> Homo sapiens

<400> 139

atggatetea tgtgeaagaa aatgaageae etgtggttet teeteetget ggtggegget 60 ceeagatggg teetgteeca getgeagetg eaggagtegg geeeaggaet aetgaageet 120 teggagaeee tgteeeteae etgeaetgte tetggegget eeateageag teetggttae 180 taeegggget ggateegeea geeeeeaggg aaggggetgg agtggattgg gagtatetat 240 aaaagtggga geaeetaeea eaaceegtee etcaaggagte gagteaeeat ateegtagae 300 aegteeaaga aeeagttete eetgaagetg agetetgtga eegeeegeaga eaeggetgtg 360 tattaetgta egagaeetgt agtaegatat tttgggtggt tegaeeeetg gggeeaggga 420 aeeetggtea eegteetee agetageaee aaggggeeat eegteteee eetggegee 480 tgeteeagga geaeeteega gageaeagee geeetggget geetggteaa ggaetaette 540 eeeggetgtee taeagteete aggaeeteae teeeteagea gegtggtae eggeeetee 660 ageaggettgg geaegaagae etaeeetge aaeggagate aeeggtggae eggeeetee 660 ageagettgg geaegaagae etaeeetge aaegtagate aeeageeeag eaacaeeaag 720

gtggacaaga gagttgagte caaatatggt eececatgee eaceatgeee ageaeetgag 780
ttegaggggg gaceateagt etteetgtte eececaaaac eeaaggacae teteatgate 840
teeeggacee etgaggteae gtgegtggtg gtggacgtga geeaggaaga eeeegggggg 960
cagtteaaet ggtacgtgaa tggegtggag gtgeataatg eeaagacaaa geegggggag 960
gageagttea acageaegta eegtggte agegteetea eegteetgea eeaggaetgg 1020
etgaaeeggea aggagtacaa gtgeaaggte teeaacaaaag geeteeegte eteeategag 1080
aaaaceatet eeaaageeaa aggeeageee eggaggeeae aggtgtacae eetgeeeea 1140
teeeaggagg agatgacaa gageggage aatggeegee eggagaacaa etacaagaee 1260
acgeeteeeg tgetggaete egaeggetee ttetteetet acageagget aacegtggae 1320
aagageaggt ggeaggagg gaatgtette teatgeteeg tgatgeatga ggetetgeae 1380
aaceaetaca eacagaagag eeteteeetg teeteggta aatga

<210> 140

<211> 474

<212> PRT

<213> Homo sapiens

<400> 140

Met Asp Leu Met Cys Lys Lys Met Lys His Leu Trp Phe Phe Leu Leu

1 5 10 15

Leu Val Ala Ala Pro Arg Trp Val Leu Ser Gln Leu Gln Leu Gln Glu
20 25 30

Ser Gly Pro Gly Leu Leu Lys Pro Ser Glu Thr Leu Ser Leu Thr Cys
35 40 45

Thr Val Ser Gly Gly Ser Ile Ser Ser Pro Gly Tyr Tyr Gly Gly Trp 82/88

Ile Arg Gln Pro Pro Gly Lys Gly Leu Glu Trp Ile Gly Ser Ile Tyr 65 70 75 80

Lys Ser Gly Ser Thr Tyr His Asn Pro Ser Leu Lys Ser Arg Val Thr

85 90 95

Ile Ser Val Asp Thr Ser Lys Asn Gln Phe Ser Leu Lys Leu Ser Ser

100 105 110

Val Thr Ala Ala Asp Thr Ala Val Tyr Tyr Cys Thr Arg Pro Val Val
115 120 125

Arg Tyr Phe Gly Trp Phe Asp Pro Trp Gly Gln Gly Thr Leu Val Thr
130 135 140

Val Ser Ser Ala Ser Thr Lys Gly Pro Ser Val Phe Pro Leu Ala Pro 145 150 155 . 160

Cys Ser Arg Ser Thr Ser Glu Ser Thr Ala Ala Leu Gly Cys Leu Val 165 170 175

Lys Asp Tyr Phe Pro Glu Pro Val Thr Val Ser Trp Asn Ser Gly Ala 180 185 190

Leu Thr Ser Gly Val His Thr Phe Pro Ala Val Leu Gln Ser Ser Gly
195 200 205

Leu Tyr Ser Leu Ser Ser Val Val Thr Val Pro Ser Ser Ser Leu Gly 83/88

Thr Lys Thr Tyr Thr Cys Asn Val Asp His Lys Pro Ser Asn Thr Lys 225 230 235 240

220

Val Asp Lys Arg Val Glu Ser Lys Tyr Gly Pro Pro Cys Pro Pro Cys
245
250
255

Pro Ala Pro Glu Phe Glu Gly Gly Pro Ser Val Phe Leu Phe Pro Pro 260 265 270

Lys Pro Lys Asp Thr Leu Met Ile Ser Arg Thr Pro Glu Val Thr Cys
275
280
285

Val Val Val Asp Val Ser Gln Glu Asp Pro Glu Val Gln Phe Asn Trp 290 295 300

Tyr Val Asp Gly Val Glu Val His Asn Ala Lys Thr Lys Pro Arg Glu 305 310 315 320

Glu Gln Phe Asn Ser Thr Tyr Arg Val Val Ser Val Leu Thr Val Leu
325 330 335

His Gln Asp Trp Leu Asn Gly Lys Glu Tyr Lys Cys Lys Val Ser Asn 340 345 350

Lys Gly Leu Pro Ser Ser Ile Glu Lys Thr Ile Ser Lys Ala Lys Gly
355 360 365

Gln Pro Arg Glu Pro Gln Val Tyr Thr Leu Pro Pro Ser Gln Glu Glu 370 375 380

Met Thr Lys Asn Gln Val Ser Leu Thr Cys Leu Val Lys Gly Phe Tyr 385 390 395 400

Pro Ser Asp Ile Ala Val Glu Trp Glu Ser Asn Gly Gln Pro Glu Asn 405 410 415

Asn Tyr Lys Thr Thr Pro Pro Val Leu Asp Ser Asp Gly Ser Phe Phe
420 425 430

Leu Tyr Ser Arg Leu Thr Val Asp Lys Ser Arg Trp Gln Glu Gly Asn
435
440
445

Val Phe Ser Cys Ser Val Met His Glu Ala Leu His Asn His Tyr Thr 450 455 460

Gln Lys Ser Leu Ser Leu Ser Leu Gly Lys 465 470

<210> 141

<211> 708

<212> DNA

<213> Homo sapiens

<400> 141

atggacatga gggtccccgc tcagctcctg gggcttctgc tgctctggct cccaggtgcc 60 agatgtgcca tccagttgac ccagtctcca tcctccctgt ctgcatctgt aggagacaga 120

gtcaccatca cttgccggc aagtcaggc attagcagt ctttagcctg gtatcagcag 180
aaaccagga aagctcctaa gctcctgatc tatgatgcct ccaatttgga aagtggggtc 240
ccatcaaggt tcagcggcag tggatctggg acagatttca ctctcaccat cagcagcctg 300
cagcctgaag attttgcaac ttattactgt caacagttta atagttaccc gacgttcggc 360
caagggacca aggtggaaat caaacgtacg gtggctgcac catctgtctt catcttcccg 420
ccatctgatg agcagttgaa atctggaact gcctctgttg tgtgcctgct gaataacttc 480
tatcccagag aggccaaagt acagtggaag gtggataacg ccctccaatc gggtaactcc 540
caggagagtg tcacagagca ggacagcaag gacagcacct acagcctcag cagcacctg 600
acgctgagca aagcagacta cgagaaacac aaagtctacg cctgcgaagt cacccatcag 660
ggcctgagct cgcccgtcac aaagagcttc aacaggggag agtgttga 708

<210> 142

<211> 235

<212> PRT

<213> Homo sapiens

<400> 142

Met Asp Met Arg Val Pro Ala Gln Leu Leu Gly Leu Leu Leu Trp

1 5 10 15

Leu Pro Gly Ala Arg Cys Ala Ile Gln Leu Thr Gln Ser Pro Ser Ser
20 25 30

Leu Ser Ala Ser Val Gly Asp Arg Val Thr Ile Thr Cys Arg Ala Ser

35 40 45

Gln Gly Ile Ser Ser Ala Leu Ala Trp Tyr Gln Gln Lys Pro Gly Lys
50 55 60

| Ala | Pro | Lys | Leu | Leu | Ile | Tyr | Asp | Ala | Ser | Asn | Leu | Glu | Ser | Gly | Val |
|-----|-----|-----|-----|-----|-----|-----|-----|-----|-----|-----|-----|-----|-----|-----|-----|
| 65 | | | | | 70 | | | | | 75 | | | | | 80 |
| Pro | Ser | Arg | Phe | Ser | Gly | Ser | Gly | Ser | Gly | Thr | Asp | Phe | Thr | Leu | Thr |
| | | | | 85 | | | | | 90 | | | | | 95 | |
| Ile | Ser | Ser | Leu | Gln | Pro | Glu | Asp | Phe | Ala | Thr | Tyr | Tyr | Cys | Gln | Gln |
| | | | 100 | | | | | 105 | | | | | 110 | | |
| Phe | Asn | Ser | Tyr | Pro | Thr | Phe | Gly | Gln | Gly | Thr | Lys | Val | Glu | Ile | Lys |
| | | 115 | | | | | 120 | | | | | 125 | | | |
| Arg | Thr | Val | Ala | Ala | Pro | Ser | Val | Phe | Ile | Phe | Pro | Pro | Ser | Asp | Glu |
| | 130 | | | | | 135 | | | | | 140 | | | | |
| Gln | Leu | Lys | Ser | Gly | Thr | Ala | Ser | Val | Val | Cys | Leu | Leu | Asn | Asn | Phe |
| 145 | | | | | 150 | | | | | 155 | | | | | 160 |
| Tyr | Pro | Arg | Glu | Ala | Lys | Val | Gln | Trp | Lys | Val | Asp | Asn | Ala | Leu | Gln |
| | | | | 165 | | | | | 170 | | | | | 175 | |
| Ser | Gly | Asn | Ser | G1n | Glu | Ser | Val | Thr | Glu | Gln | Asp | Ser | Lys | Asp | Ser |
| | | | 180 | | | | | 185 | | | | | 190 | | |
| Thr | Tyr | Ser | Leu | Ser | Ser | Thr | Leu | Thr | Leu | Ser | Lys | Ala | Asp | Tyr | Gĺu |
| | | 195 | | | | | 200 | | | | | 205 | | | |
| Lys | His | Lys | Val | Tyr | Ala | Cys | Glu | Val | Thr | His | Gln | Gly | Leu | Ser | Ser |
| | 210 | | | | | 215 | | | | | 220 | | | | |

| Pro | Val | Thr | Lys | Ser | Phe | Asn | Arg | Gly | Glu | Cys |
|-----|-----|-----|-----|-----|-----|-----|-----|-----|-----|-----|
| 225 | | | | | 230 | | | | | 235 |